PCysMl:

5'ATAT<u>GGATGC</u>ATCGAGGGTAGGGCCGATGCCGGCTACGCCCGGCCCACCCCGGCTACCCCGGCTACGCCCCGGC

Please amend the paragraph on page 33, lines 19 through 25 as follows:

Plasmid pGS21 (see above) was used as the starting vector for cloning the deletion mutant DM1. The bp 399 -1374 fragment of the cDNA for rPh1 p 5b was amplified in a PCR using the following primers:

MP2 sense:

5'-GCTAGCCGGCGAGCTGCAGATCATCG-3' (SEQ ID NO 99)

REMARKS

The above amendment is submitted in response to the Notice of Non-Compliant Amendment mailed September 20, 2001. No new matter is introduced and it is respectfully requested that the application be examined on its merits.

Respectfully submitted,

Nancy J. Axelrod

Registration No. 44,014

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Attorney Docket No.: MERCK-2034

Filed: October 22, 2001

VERSION WITH MARKINGS TO SHOW CHANGES MADE

IN THE SPECIFICATION:

Please amend pages 6, 9-11, 28 and 31-33 as follows:

Please amend the paragraph beginning on page 5, line 26, through page 6, line 12 as follows:

Using the single-letter code for amino acids, the sequence of Phl p 5b is as follows:

ADAGYAPATPAAAGAAAGKATTEEQKLIEDINVGFKAAVAAAASVPAADK							
1 .	10	20	30	40	50		
FKTFEAAFTSSSKAAAAKAPGLVPKLDAAYSVAYKAAVGATPEAKFDSFV							
51	60	70	80	, St	100		
ASLTEALRVIAGALEVHAVKPVTEEPGMAKIPAGELQIIDKIDAAFKVAA							
101	110	120	130	140	150		
TAAATAPADDKFTVFEAAFNKAIKESTGGAYDTYKCIPSLEAAVKQAYAA							
151	160	170	180	190	200		
TVAAAPQVKYAVFEAALTKAITAMSEVQKVSQPATGAATVAAGAATTAAG							
201	210	220	230	24	0 250		
AASGAATVAAGGYKV (SEQ ID NO 87)							
251	250 255						

Please amend the paragraph on page 9, lines 1-9 as follows:

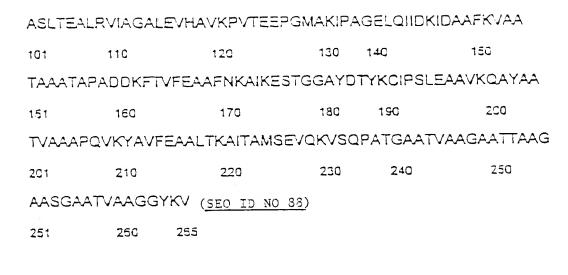
In this context, PM1 denotes point mutation 1 and has the following sequence (the amino acids which are replaced as compared with Ph1 p 5b are printed in bold):

ADAGYAPATPAAAGAAAGKATTEECKLIEDIDVGFKAAVAAAASVPAALA

1 10 20 30 40 50

FKTFEAAFTSSSKAAAAKAPGLVPKLDAAYSVAYKAAVGATPEAKFDSFV

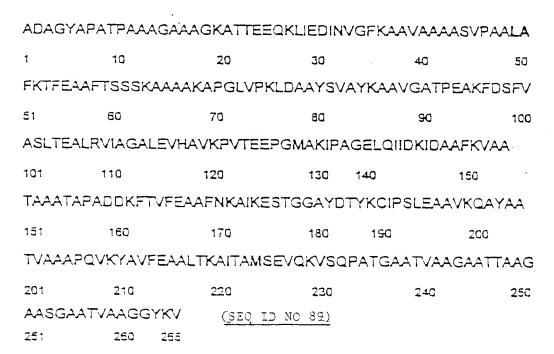
51 60 70 80 90 100



Please amend the paragraphs beginning on page 9, line 10 through page 10, line 16 as follows:

The other particularly preferred peptides have the following sequences:

PM2
$$(D^{49} \rightarrow L, K^{50} \rightarrow A)$$
:



PM3 $(A^{13} \rightarrow C)$:

ALAG	1APA (PAAC	GAAAGKATTI	EEQKLIEDIN	VGFKAAVAA	AASVPAADK
1	10	20	30	40	50
FKTFE	AAFTSSSKA	AAAKAPGLV	PKLDAAYSV	'AYKAAVGAT	TPEAKFDSFV
51	60	70	08	90	100
ASLTE	ALRVIAGAL	EVHAVKPVTE	EEPGMAKIPA	AGELQIIDKIE)AAFKVAA
101	110	120	130	140	150
TAAAT	APADDKFT	/FEAAFNKAIH	KESTGGAYD	TYKCIPSLE	VAVKQAYAA
151	160	170	180	190	200
TVAAA	(PQVKYAVF	EAALTKAITAN	NSEVQKVSC	PATGAATVA	VAGAATTAAG
201	210	220	230	240	250
AASGA	YATVAAGGY	KV (SEQ ID	NO 90)		
251	250	265			
DM1 (Δ	$K^{SO} \rightarrow P^{\Delta LS}$	12 , $D^{49} \rightarrow L$) :		
ADAGY	APATPAAAC	BAAAGKATTE	EQKLIEDINV	GFKAAVAAA	ASVPAALA
1	10	20	30	40	50
GELQII	DKIDAAFKV	AATAAATAPA	DOKFTVFEA	AFNKAIKEST	rggaydtyk
51	60	70	80	90	100
CIPSLE	AAVKQAYA	ATVAAAPQVK	YAVFEAALT	KAITAMSEV	QKVSQPATG
103	110	. 120	130	140	15G
AATVA	AGAATTAAG	SAASGAATVA	AGGYKV <u>(s</u>	EQ ID NO 91)
154	160	170	180		
		.,, -			
DM 2 (3	1 F ³¹ - G ¹⁷	3, 549 - 5,	$\mathbb{K}^{50} \rightarrow \mathbb{A}$:		
ADAGY	YAPATPAAA	GAAAGKATTE	EECKLIEDIN	/GFKAAVAA	aasvpaa la
1	10	20	3G	40	50
GAYDI	TYKCIPSLEA	AVKQAYAAT	VAAAPQVKY	AVFEAALTK	AITAMSEVQK
51	6 0	70	8C	9 0	100
VSQPA	TGAATVAA	GAATTAAGAA	KSGAATVAA	GGYKV (sa	מו מו מו פו
102	110	120		137	4 40 21 /

Please amend the paragraph on page 11, lines 2 through lines 12 as follows:

This sequence corresponds to that of DM2 where, however, the amino acids of positions 179-217 of the starting peptide Ph1 p 5b additionally exhibit an altered sequence and all the subsequent amino acids are missing.

DM3 (
$$\Delta$$
 A¹⁵⁴ - T¹⁷⁷, A²²⁰ \rightarrow T):

ADAGYAPATPAAAGAAAGKATTEEQKLIEDINVGFKAAVAAAASVPAADK FKTFEAAFTSSSKAAAAKAPGLVPKLDAAYSVAYKAAVGATPEAKFDSFV ASLTEALRVIAGALEVHAVKPVTEEPGMAKIPAGELQIIDKIDAAFKVAA TAAGGAYDTYKCIPSLEAAVKQAYAATVAAAPQVKYAVFEAALTKTITAMS EVOKVSQPATGAATVAAGAATTAAGAASGAATVAAGGYKV (SEQ ID NO 93)

Please amend the paragraph on page 28, lines 3 through lines 48 as follows:

Tab. 1: Dodecapeptides which are based n the Ph1 p 5b sequence and which are used for determining the T cell-reactive regions

1	ACAGYAPATPAA		4-4	KIPAGELQIIDK	(SEQ ID NO 44)
2	GYAPATPAAAGA	(SEQ ID NO I)	45	AGELQIIDKIDA	(SEQ ID NO 45)
3	PATPAAAGAAAG	(SEQ ID NO 2)	46	LOHDKIDAAFK	(SEQ ID NO 46)
4	PAAAGAAAGKAT	(SEQ ID NO 3)	47	IDKIDAAFKVAA	(SEQ ID NO 47)
5	AGAAAGKATTEE	(SEQ ID NO 4)	48	IDAAFKVAATAA	(SEQ ID NO 48)
â	AAGKATTEEQKL	(SEQ ID NO 5)	49	AFKVAATAAATA	(SEQ ID NO 49)
7	KATTEEOKLIED	(SEQ ID NO 6)	50	VAATAAATAPAD	(SEQ ID NO 50)
В	TEECKLIEDINV	(SEQ ID NO 7)	51	TAAATAPADDKF	(SEQ ID NO 51)
9	QKLIEDINVGFK	(SEQ ID NO 8)	52	ATAPADCKFT/F	(SEQ ID NO 52)
10	IEDINVGFKAAV	(SEQ ID NO 9)	53	PADDKFTVFEAA	(SEQ ID NO 53)
11	INVGFKAAVAAA	(SEQ ID NO 10)	54	DKFTVFEAAFNK	(SEQ ID NO 54)
12	GFKAAVAAAASV	(SEQ ID NO 11)	55	TVFEAAFNKAIK	
13	AAVAAAASVPAA	(SEQ ID NO 12)	56	EAAFNKAIKEST	(SEQ ID NO 55)
	AAAASVPAADKF	(SEQ ID NO 13)	57	FNKAIKESTGGA	(SEQ ID NO 56)
14 15	ASVPAADKFKTF	(SEQ ID NO 14)	58	AIKESTGGAYDT	(SEQ ID NO 57)
	PAADKFKTFEAA	(SEQ ID NO 15)	59	ESTGGAYDTYKC	(SEQ ID NO 58)
16		(SEQ ID NO 16)	60	GGAYDTYKCIPS	(SEQ ID NO 59)
17	DKFKTFEAAFTS	(SEQ ID NO 17)	61	YDTYKCIPSLEA	(SEQ ID NO 60)
18	KTFEAAFTSSSK	(SEQ ID NO 18)	62	YKCIPSLEAAVK	(SEQ ID NO 61)
19	EAAFTSSSKAAA	(SEQ ID NO 19)	63	IPSLEAAVKOAY	(SEQ ID NO 62)
20	FTSSSKAAAAKA	(SEQ ID NO 20)	64	LEAAVKOAYAAT	(SEQ ID NO 63)
21	SSKAAAAKAPGL	(SEQ ID NO 21)	65	AVKQYAATYAA	(SEQ ID NO 64)
22	AAAAKAPGLVPK	(SEQ ID NO 22)		CAYAATVAAAPO	(SEQ ID NO 65)
23	AKAPGLVPKLDA	(SEQ ID NO 23)	66		(SEQ ID NO 66)
24	PGLVPKLDAAYS	(SEQ ID NO 24)	67	AATVAAAPOVKY	(SEQ ID NO 67)
25	VPKLDAAYSVAY	(SEQ ID NO 25)	83	VAAAPQVKYAVF	(SEQ ID NO 68)
25	LDAAYSVAYKAA	(SEQ ID NO 26)	69	APQVKYAVFEAA	(SEQ ID NO 69)
27	AYSVAYKAAVGA	(SEQ ID NO 27)	70	VKYAVFEAALTK	(SEQ ID NO 70)
28	VAYKAAVGATPE	(SEQ ID NO 28)	71	AVFEAALTKAIT	(SEQ ID NO 71)
29	KAAVGATPEAKF	(SEQ ID NO 29)	72	EAALTKAITAMS	(SEQ ID NO 72)
30	VGATPEAKFDSF	(SEQ ID NO 30)	73	LTKAITAMSEVQ	(SEQ ID NO 73)
31	TPEAKFOSFVAS	(SEQ ID NO 31)	74	AITAMSEVQKVS	(SEQ ID NO 74)
32	AKFDSFVASLTE	(SEQ ID NO 32)	75	AMSEVQKVSQPA	(SEQ ID NO 75)
33	DSFVASLTEALR	(SEQ ID NO 33)	76	EVOKVSOPATGA	(SEQ ID NO 76)
34	VASLTEALRVIA	(SEQ ID NO 34)	77	KVSQPATGAATV	(SEQ ID NO 77)
35	LTEALRVIAGAL	(SEQ ID NO 35)	73	QPATGAATVAAG	(SEQ ID NO 78)
36	ALRVIAGALEVH	(SEQ ID NO 36)	79	TGAATVAAGAAT	(SEQ ID NO 79)
37	VIAGALEVHAVK	(SEQ ID NO 37)	80	ATVAAGAATTAA	(SEQ ID NO 80)
38	GALEVHAVKPVT	(SEQ ID NO 38)	81	AAGAATTAAGAA	(SEQ ID NO 81)
39	EVHAVKPVTEEP	(SEQ ID NO 39)	82	AATTAAGAASGA	(SEQ ID NO 82)
40	AVKPVTEEPGMA	(SEQ ID NO 40)	83	TAAGAASGAATV	(SEQ ID NO 33)
41	PYTEEPGMAKIP	(SEQ ID NO 41)	84	GAASGAATVAAG	(SEQ ID NO 84)
42	EEPGMAKIPAGE	(SEQ ID NO 42)	85	SGAATVAAGGYX	(SEQ ID NO 85)
43	GMAKIPAGELOI	(SEQ ID NO 43)	86	GAATVAAGGYKV	(SEQ ID NO 86)
~ 3	GMAIN AGELDI	(334,25.15.15)			

Please amend the paragraph on page 31, lines 35 through page 32, lines 7 as follows:

Fragment 1:

Ph1 p 5b sense:

5'-ATAT<u>GGATOC</u>ATCGAGGGAAGGGCCGATGCCGGCTACGCC-3' (SEQ ID NO 94)

```
MP1 antisense:

5'-GAACGCTAGGGCCGCAGGGACGCTGGC-3' (SEQ ID NO 95)

Fragmant 2:

MP1 sense:

5'-GCGCTAGCGTTCAAGACCTTCGAG-3' (SEQ ID NO 96)

Ph1 p 5b antisense:

5'-ATATAAGCTTTCCTCTGAAGGGAAGGCAACCC-3' (SEQ ID NO 97)
```

Please amend the paragraph on page 32, lines 30-38 as follows:

The point mutant rPh1 p 5b PM1 was prepared in analogy with PM2. It contains, as the result of a PCR error, an additional point mutation: $N^{32} \rightarrow D$.

In order to clone this point mutant, the entire cDNA for rPh1 p 5b in vector pGS13 was amplified in a PCR using the following primers.

```
PCysMl:
5'ATAT<u>GGATCC</u>ATCGAGGGTAGGGCCGATGCCGGGCTACGCCCGGGC
CACCCCGGCT<u>GCATGC</u>GGAGCG-3' (SEQ ID NO 98)
```

Please amend the paragraph on page 33, lines 19 through 25 as follows:

Plasmid pGS21 (see above) was used as the starting vector for cloning the deletion mutant DM1. The bp 399 -1374 fragment of the cDNA for rPh1 p 5b was amplified in a PCR using the following primers:

```
MP2 sense:
5'-GCTAGCCGGCGAGCTGCAGATCATCG-3' (SEQ ID NO 99)
```